

**SUBSTITUTE COMBINED DECLARATION AND POWER OF ATTORNEY**

As a below named inventor, I hereby declare that:

My residence, post office address and citizenship are as stated below next to my name; and

I verily believe that I am the original, first and sole inventor (if only one name is listed below) or an original, first and joint inventor (if plural names are listed below) of the subject matter which is claimed and for which a patent is sought on the invention entitled: **PEPTIDE NUCLEIC ACID CONJUGATES** the specification of which was filed on October 6, 1995 as International Application Serial No. PCT/US95/12931.

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<u>Denmark</u>	<u>987/91</u>	<u>May 24, 1991</u>	<u>Yes</u>
<u>Denmark</u>	<u>510/92</u>	<u>April 15, 1992</u>	<u>Yes</u>

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<u>08/088,658</u>	<u>July 2, 1993</u>	<u>Patented</u>
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<u>08/275,951</u>	<u>July 15, 1994</u>	<u>Pending</u>

I hereby appoint the following attorney(s) and/or agent(s) to prosecute this application and to transact all business in the Patent and Trademark Office connected therewith: **John W. Caldwell** and **Michael P. Straher**, Registration Nos. 28,937 and 38,325 of the firm of **WOODCOCK WASHBURN KURTZ MACKIEWICZ & NORRIS LLP**, One Liberty Place - 46th Floor, Philadelphia, Pennsylvania 19103, and **Herb Boswell** and **Laurel Bernstein**, Registration Nos. 27,311 and 37,280, of **ISIS Pharmaceuticals, Inc.**, 2292 Faraday Avenue, Carlsbad, California 92008.

Address all telephone calls and correspondence to:

Michael P. Straher
WOODCOCK WASHBURN KURTZ MACKIEWICZ & NORRIS LLP
One Liberty Place - 46th Floor
Philadelphia, PA 19103
Telephone No. 215-568-3100.

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

1	Full Name: Peter NIELSEN	Inventor's Signature:	Date:
	Residence: Kokkedal, DENMARK	Citizenship: Denmark	
	Post Office Address: Hjortevanget 509, DK 2980 Kokkedal, DENMARK		
2	Full Name: Ole BUCHARDT (deceased)	Inventor's Signature: <i>D. Buchardt</i> Mrs. Dorte BUCHARDT Signing upon information and belief as the Representative of the Estate of Ole BUCHARDT	Date 5/299
	Residence: Vaerlose, DENMARK	Citizenship: Denmark	
	Post Office Address: Sondergardsvej 73, DK 3500, Vaerlose, DENMARK		
3	Full Name: Soren Holst SONNICHSEN	Inventor's Signature:	Date
	Residence: Tastrup, DENMARK	Citizenship: Denmark	
	Post Office Address: Gronhojgardsvej 13, DK 2630 Tastrup, DENMARK		
4	Full Name: Jesper LOHSE	Inventor's Signature:	Date
	Residence: Copenhagen N, DENMARK	Citizenship: Denmark	
	Post Office Address: Staerevej 52, 2.tv, DK 2400, Copenhagen N, DENMARK		
5	Full Name: Michael EGHOLM	Inventor's Signature:	Date
	Residence: Wayland, Massachusetts	Citizenship: Denmark	
	Post Office Address: 34 Grove Street, Wayland, Massachusetts 01778		

6	Full Name: Muthiah MANOHARAN	Inventor's Signature:	Date:
	Residence: Carlsbad, California	Citizenship: United States	
	Post Office Address: 7634 Reposado Drive, Carlsbad, California 92009		
7	Full Name: John KIELY	Inventor's Signature:	Date
	Residence: San Diego, California	Citizenship: United States	
	Post Office Address: 4230 Corte Facil, San Diego, California 92130		
8	Full Name: Michael GRIFFITH	Inventor's Signature:	Date
	Residence: San Diego, California	Citizenship: United States	
	Post Office Address: 5676 Greenshade Road, San Diego, California 92121		
9	Full Name: Kelly SPRANKLE	Inventor's Signature:	Date
	Residence: Vista, California	Citizenship: United States	
	Post Office Address: 920 Sycamore Avenue, Apt. 61, Vista, California 92083		
	Full Name:	Inventor's Signature:	Date
	Residence:	Citizenship:	
	Post Office Address:		



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		Mrs. Dorte BUCHARDT Signing upon information and belief as the Representative of the Estate of Ole BUCHARDT	
	Residence: Vaerlose, DENMARK	Citizenship: Denmark	
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4	Full Name: Jesper LOHSE	Inventor's Signature:	Date
	Residence: Copenhagen N, DENMARK	Citizenship: Denmark	
	Post Office Address: Staerevej 52, 2.tv, DK 2400, Copenhagen N, DENMARK		
5	Full Name: Michael EGHOLM	Inventor's Signature:	Date
	Residence: Lexington, Massachusetts	Citizenship: Denmark	
	Post Office Address: 1231 Lexington Ridge Drive, Lexington, Massachusetts 02173		

6	Full Name: Muthiah MANOHARAN	Inventor's Signature: <i>Muthiah Manohan</i>	Date: 4/16/99
	Residence: Carlsbad, California	Citizenship: United States	
	Post Office Address: 7634 Reposado Drive, Carlsbad, California 92009		
7	Full Name: John KIELY	Inventor's Signature: <i>John S Kiely</i>	Date: 4/12/99
	Residence: San Diego, California	Citizenship: United States	
	Post Office Address: 4230 Corte Facil, San Diego, California 92130		
8	Full Name: Michael GRIFFITH	Inventor's Signature: <i>Michael Griffith</i>	Date: 4/12/99
	Residence: San Diego, California	Citizenship: United States	
	Post Office Address: 5676 Greenshade Road, San Diego, California 92121		
9	Full Name: Kelly SPRANKLE	Inventor's Signature: <i>Kelly Sprankle</i>	Date: 4/16/99
	Residence: Vista, California	Citizenship: United States	
	Post Office Address: 920 Sycamore Avenue, Apt. 61, Vista, California 92083		
	Full Name:	Inventor's Signature:	Date
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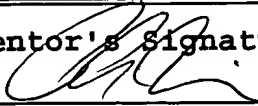

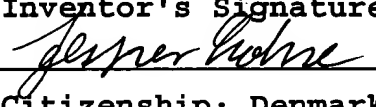
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	Post Office Address: Sondergardsvej 73, DK 3500, Vaerloose, DENMARK		
3	Full Name: Soren Holst SONNICHSEN	Inventor's Signature: 	Date: 18/2-99
	Residence: Tastrup, DENMARK	Citizenship: Denmark	
	Post Office Address: Gronhojgardsvej 13, DK 2630 Tastrup, DENMARK		
4	Full Name: Jesper LOHSE	Inventor's Signature: 	Date: 22/2 99
	Residence: Copenhagen N, DENMARK	Citizenship: Denmark	
	Post Office Address: Staerevej 52, 2.tv, DK 2400, Copenhagen N, DENMARK		
5	Full Name: Michael EGHOLM	Inventor's Signature:	Date
	Residence: Wayland, Massachusetts	Citizenship: Denmark	
	Post Office Address: 34 Grove Street, Wayland, Massachusetts 01778		

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DOCKET NO.: ISIS-2112
PATENT APPLICATION



SERIAL NO.: 08/817,067
FILED: April 4, 1997

08/1634

#8
B. Webb
6/9/99

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

RECEIVED

In re application of:

Nielsen et al.

MAY 05 1999

Serial No.: 08/817,067

Group Art Unit: 1634
MATRIX CUSTOMER
SERVICE CENTER

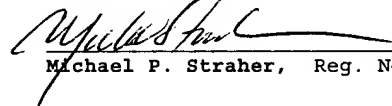
Filed: April 4, 1997

Examiner: A. Marschel

For: PEPTIDE NUCLEIC ACID CONJUGATES

I, Michael P. Straher, Registration No. 38,325 certify that this correspondence is being deposited with the U.S. Postal Service as First Class mail in an envelope addressed to the Assistant Commissioner for Patents, Washington, D.C. 20231.

On April 28, 1999


Michael P. Straher, Reg. No. 38,325

Assistant Commissioner
for Patents,
Washington, D.C. 20231

Dear Sir:

TRANSMITTAL OF SUBSTITUTE COMBINED DECLARATION
AND POWER OF ATTORNEY

Transmitted herewith are four (4) executed Substitute Combined Declaration and Power of Attorney forms in the above-identified patent application. The subject application is a U.S. National Phase filing of International Application S.N. PCT/US95/12931, filed October 6, 1995, which is referenced in the substitute declarations.

DOCKET NO.: ISIS-2112
PATENT APPLICATION


SERIAL NO.: 08/817,067
FILED: April 4, 1997

Applicants are filing the substitute declarations due to a clerical error on the previously filed declarations in which one of the priority applications was listed twice (08/088,658) and and another priority application was omitted (08/088,661).

(XX) The Commissioner is hereby authorized to charge payment of any fees associated with this communication or credit any overpayment to Deposit Account No. 23-3050.

This sheet is attached in duplicate.

Respectfully submitted,


Michael P. Straher
Registration No. 38,325

Date: April 28, 1999

WOODCOCK WASHBURN KURTZ
MACKIEWICZ & NORRIS LLP
One Liberty Place - 46th Floor
Philadelphia, PA 19103
(215) 568-3100